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APPENDIX

Changes made to the specification are shown below, with insertions underlined and deletions bracketed.

RNA was isolated from CLL or whole mononuclear cells using Ultraspec reagent (Biotecx, Houston, TX). cDNA was synthesized from 10 ug of total RNA using oligo d(T) primers and Maloney murine leukemia virus reverse transcriptase in a final volume of 40 uL (Stratagene, La Jolla, CA). One mL of the first strand cDNA product was then used as template for PCR amplification with AmpliTaq DNA polymerase (Roche Molecular Systems, Branchburg, NJ) by 40 thermocycles of 94°C for 1 minute, 60°C for 1 minute and 72°C for 1 minute. The PDE PCR assay products were as follows with oligonucleotide sequences given 5'->3': Human PDE1B1 (Genbank accession # U56976) was 430 bp (1st base 1660; sense = GTC TTC ATT GAG TCC AAA GTG (SEQ ID NO.:1), antisense = GAC CTG CCA GCT AAG ATC TGG (SEQ ID NO.:2)). Human PDE3A (cGIP1, HSPDE3B) (X95520) was 340 bp (1st base 2999, sense = GTA ACT CCT ATG ATG CTG CTG G (SEQ ID NO.:3), antisense = CTA TTC CTC TTC ATC TGC CTC (SEQ ID NO.:4)). Of note, these PDE3 PCR oligonucleotides are selective for the human cGIP1 PDE , homologous to rat PDE3A, as the amplified sequence has only 50% nucleotide homology to the cardiac/platelet form of human PDE3 (cGIP2). Human PDE4A (M37744) was 461 bp (1st 1819, sense = GGA GGA AGA AAT ATC AAT GGC CC (SEQ ID NO.:5), antisense = GAT GTG TCC TCC CCA AAT GTC (SEQ ID NO.:6)). Human PDE4B (L20966) was 479 bp (1st bp 2213, sense = ATT CTG AAG GAC CTG AGA AGG (SEQ ID NO.:7), antisense = CAG TGA GTT CAG TCA CTG TCG (SEQ ID NO.:8)). For hybridization to Northern blots, these PCR products were subcloned into a plasmid vector

(pCRII, Invitrogen, Carlsbad, CA) and subsequently utilized for PCR-based amplification of a32P dATP-labelled probes.